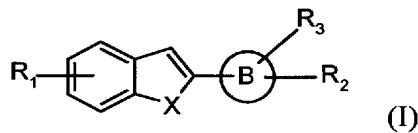


Amendments to Claims

The listing of claims will replace all prior versions, and listings of claims in the application:

Listing of Claims:

1. (Currently Amended): A method of treating or inhibiting a disorders associated with the activation of large conductance calcium activated potassium channels, wherein the disorder is selected from the group consisting of: urinary incontinence, overactive bladder, pollakiuria, urge incontinence, diseases associated with detrusor instability, irritable bladder, cystitis, urethritis, and kidney stone ailments, which comprises administering to a subject in need thereof an effective amount of a compound according to formula (I):



wherein:

R_1 is absent or represents up to three substituents independently selected from (C₁₋₆)alkyl, (C₂₋₆)alkenyl, (C₃₋₆)cycloalkyl, aryl, (C₁₋₆)alkyl-aryl, ~~heterocycle, (C₁₋₆)alkyl-heterocycle, OR_a, SR_a~~, hydroxy, halogen, nitro, trifluoromethyl, cyano, COR_a, CO₂R_a, SO₃H, (C₁₋₆)alkyl-CO₂-(C₁₋₆)alkyl, CONR_aR_b, and NR_aR_b;

where each said (C₁₋₆)alkyl, (C₂₋₆)alkenyl, and (C₃₋₆)cycloalkyl group is unsubstituted or substituted with 1 to 5 substituents selected from the group consisting of: halo, -OR', -SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R';

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where each said aryl group is unsubstituted or substituted with 1 to 3 substituents selected from the group consisting of: halo, -OR', -SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R'; and

~~where each said heterocycle group is unsubstituted or substituted with 1 to 3 substituents selected from the group consisting of: halo, OR', SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, N(R')₂, CH₂N(R')₂, nitro, cyano, CO₂R', CON(R')₂, COR', and -NR'C(O)R';~~

each R' is independently H or unsubstituted (C₁₋₆)alkyl;

X is NR_a, O, or S;

B is aryl or heterocycle phenyl;

R₂ is absent or represents up to three substituents independently selected from (C₁₋₆)alkyl, (C₂₋₆)alkenyl, (C₃₋₆)cycloalkyl, aryl, (C₁₋₆)alkyl-aryl, heterocycle, (C₁₋₆)alkyl-heterocycle, OR_a, SR_a, hydroxy, halogen, nitro, cyano, COR_a, CO₂R_a, SO₃H, (C₁₋₆)alkyl-CO₂-(C₁₋₆)alkyl, CONR_aR_b, and NR_aR_b;

where each said (C₁₋₆)alkyl, (C₂₋₆)alkenyl, and (C₃₋₆)cycloalkyl group is unsubstituted or substituted with 1 to 5 substituents selected from the group consisting of: halo, -OR', -SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R';

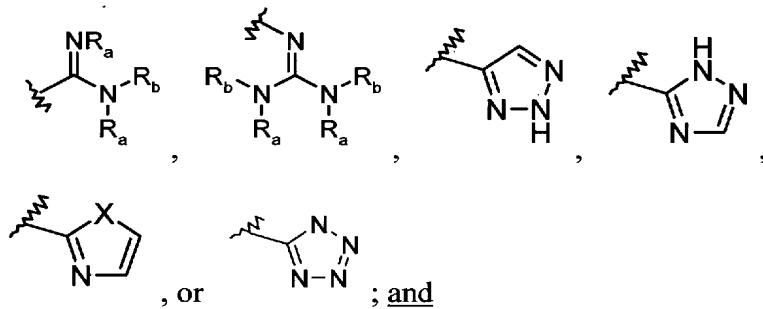
where each said aryl group is unsubstituted or substituted with 1 to 3 substituents selected from the group consisting of: halo, -OR', -SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R'; and

~~where each said heterocycle group is unsubstituted or substituted with 1 to 3 substituents selected from the group consisting of: halo, OR', SR', (C₁₋₆)alkylsulfonyl,~~

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~~(C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R';~~

R₃ is COOH, CONR_aR_b, SO₃H, SO₂NR_aR_b, CONR_aSO₂R_b.



each R_a and R_b is independently selected from hydrogen, (C₁₋₆)alkyl, aryl, and heterocycle, (C₁₋₆)alkyl-aryl, and (C₁₋₆)alkyl heterocycle;

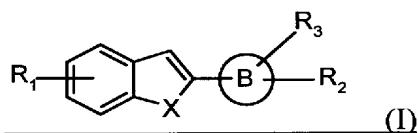
where each said (C₁₋₆)alkyl group is unsubstituted or substituted with 1 to 5 substituents selected from the group consisting of: halo, -OR', -SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R';

where each said aryl group is unsubstituted or substituted with 1 to 3 substituents selected from the group consisting of: halo, -OR', -SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R'; and

~~where each said heterocycle group is unsubstituted or substituted with 1 to 3 substituents selected from the group consisting of: halo, -OR', -SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R';~~

or a pharmaceutically acceptable salt thereof.

2. (Currently Amended): A method ~~according to claim 1~~ of relaxing bladder smooth muscle tissue through the activation of large conductance calcium activated potassium channels, which comprises administering to a subject in need thereof an effective amount of a compound according to formula (I):



wherein:

R₁ is absent or represents up to three substituents independently selected from (C₁₋₆)alkyl, (C₂₋₆)alkenyl, (C₃₋₆)cycloalkyl, aryl, (C₁₋₆)alkyl-aryl, OR_a, SR_a, hydroxy, halogen, nitro, trifluoromethyl, cyano, COR_a, CO₂R_a, SO₃H, (C₁₋₆)alkyl-CO₂(C₁₋₆)alkyl, CONR_aR_b, and NR_aR_b;

where each said (C₁₋₆)alkyl, (C₂₋₆)alkenyl, and (C₃₋₆)cycloalkyl group is unsubstituted or substituted with 1 to 5 substituents selected from the group consisting of: halo, -OR', -SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R';

where each said aryl group is unsubstituted or substituted with 1 to 3 substituents selected from the group consisting of: halo, -OR', -SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R';

each R' is independently H or unsubstituted (C₁₋₆)alkyl;

X is NR_a;

B is phenyl;

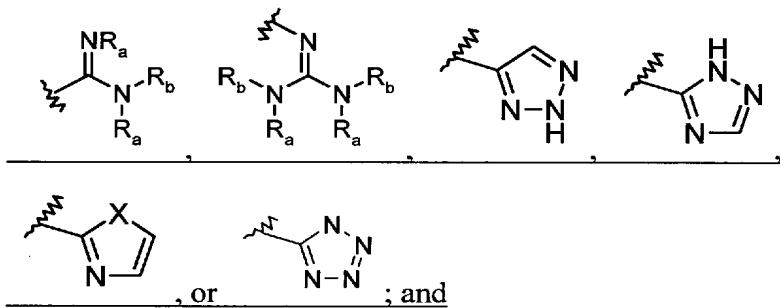
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R₂ is absent or represents up to three substituents independently selected from (C₁-6)alkyl, (C₂-6)alkenyl, (C₃-6)cycloalkyl, aryl, (C₁-6)alkyl-aryl, OR_a, SR_a, hydroxy, halogen, nitro, cyano, COR_a, CO₂R_a, SO₃H, (C₁-6)alkyl-CO₂-(C₁-6)alkyl, CONR_aR_b, and NR_aR_b:

where each said (C₁-6)alkyl, (C₂-6)alkenyl, and (C₃-6)cycloalkyl group is unsubstituted or substituted with 1 to 5 substituents selected from the group consisting of: halo, -OR', -SR', (C₁-6)alkylsulfonyl, (C₁-6)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R':

where each said aryl group is unsubstituted or substituted with 1 to 3 substituents selected from the group consisting of: halo, -OR', -SR', (C₁-6)alkylsulfonyl, (C₁-6)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R':

R₃ is COOH, CONR_aR_b, SO₃H, SO₂NR_aR_b, CONR_aSO₂R_b,



each R_a and R_b is independently selected from hydrogen, (C₁-6)alkyl, aryl, and (C₁-6)alkyl-aryl;

where each said (C₁-6)alkyl group is unsubstituted or substituted with 1 to 5 substituents selected from the group consisting of: halo, -OR', -SR', (C₁-6)alkylsulfonyl, (C₁-6)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R':

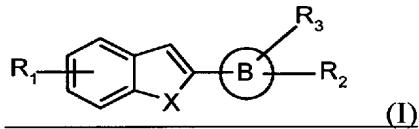
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where each said aryl group is unsubstituted or substituted with 1 to 3 substituents selected from the group consisting of: halo, -OR', -SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R';

or a pharmaceutically acceptable salt thereof.

3. (Cancelled)

4. (Currently Amended): A pharmaceutical composition which comprises a compound according to claim 1 formula (I):



wherein:

R₁ is absent or represents up to three substituents independently selected from (C₁₋₆)alkyl, (C₂₋₆)alkenyl, (C₃₋₆)cycloalkyl, aryl, (C₁₋₆)alkyl-aryl, OR_a, SR_a, hydroxy, halogen, nitro, trifluoromethyl, cyano, COR_a, CO₂R_a, SO₃H, (C₁₋₆)alkyl-CO₂-(C₁₋₆)alkyl, CONR_aR_b, and NR_aR_b;

where each said (C₁₋₆)alkyl, (C₂₋₆)alkenyl, and (C₃₋₆)cycloalkyl group is unsubstituted or substituted with 1 to 5 substituents selected from the group consisting of: halo, -OR', -SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R';

where each said aryl group is unsubstituted or substituted with 1 to 3 substituents selected from the group consisting of: halo, -OR', -SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R';

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each R' is independently H or unsubstituted (C₁₋₆)alkyl;

X is NR_a:

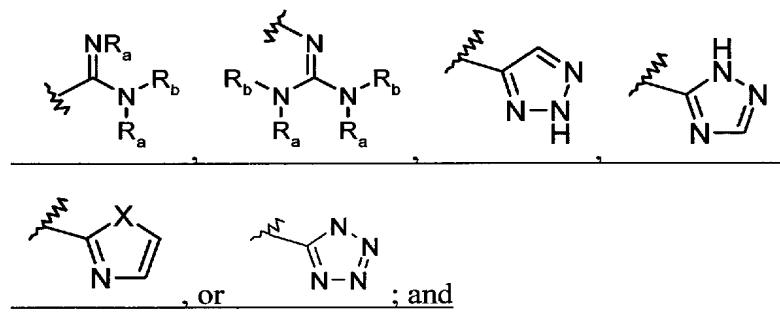
B is phenyl:

R₂ is absent or represents up to three substituents independently selected from (C₁₋₆)alkyl, (C₂₋₆)alkenyl, (C₃₋₆)cycloalkyl, aryl, (C₁₋₆)alkyl-aryl, OR_a, SR_a, hydroxy, halogen, nitro, cyano, COR_a, CO₂R_a, SO₃H, (C₁₋₆)alkyl-CO₂-(C₁₋₆)alkyl, CONR_aR_b, and NR_aR_b:

where each said (C₁₋₆)alkyl, (C₂₋₆)alkenyl, and (C₃₋₆)cycloalkyl group is unsubstituted or substituted with 1 to 5 substituents selected from the group consisting of: halo, -OR', -SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R';

where each said aryl group is unsubstituted or substituted with 1 to 3 substituents selected from the group consisting of: halo, -OR', -SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R';

R₃ is COOH, CONR_aR_b, SO₃H, SO₂NR_aR_b, CONR_aSO₂R_b,



each R_a and R_b is independently selected from hydrogen, (C₁₋₆)alkyl, aryl, and (C₁₋₆)alkyl-aryl;

where each said (C₁₋₆)alkyl group is unsubstituted or substituted with 1 to 5 substituents selected from the group consisting of: halo, -OR', -SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R';

where each said aryl group is unsubstituted or substituted with 1 to 3 substituents selected from the group consisting of: halo, -OR', -SR', (C₁₋₆)alkylsulfonyl, (C₁₋₆)alkylsulfoxyl, -N(R')₂, -CH₂N(R')₂, nitro, cyano, -CO₂R', -CON(R')₂, -COR', and -NR'C(O)R';

or a pharmaceutically acceptable salt thereof and pharmaceutically acceptable carrier.

5-8. (Withdrawn)

9. (New): The method according to claim 1 wherein the disorder is urinary incontinence.

10. (New): The method according to claim 1 wherein the disorder is an overactive bladder.

11. (New): The method according to claim 1 wherein the disorder is pollakiuria.

12. (New): The method according to claim 1 wherein the disorder is urge incontinence.

13. (New): The method according to claim 1 wherein the disorder is irritable bladder.

14. (New): The method according to claim 1 wherein the disorder is cystitis.

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15. (New): The method according to claim 1 wherein the disorder is urethritis.